

## I. AMENDMENT

### In the Claims:

Please cancel claims 1-122 without prejudice and disclaimer and add the following new claims:

- Rule 1.125*  
*B1*
- <sup>122</sup>~~123~~. (New) A method for preparing a library of regulatory DNA sequences from a cell, the method comprising:
- (a) providing a cell nucleus, wherein the nucleus comprises cellular chromatin;
  - (b) contacting the nucleus with a first enzyme, wherein the first enzyme reacts with accessible regions of cellular chromatin;
  - (c) deproteinizing the cellular chromatin to generate deproteinized DNA;
  - (d) contacting the deproteinized DNA with a second enzyme to generate DNA fragments;
  - (e) contacting the DNA fragments with a population of vector molecules, wherein the vector molecules comprise a first end that is compatible with the first enzyme and a second end that is compatible with the second enzyme, under conditions favorable to ligation of compatible ends; and
  - (f) selecting polynucleotides comprising a DNA fragment ligated to a vector molecule.

<sup>123</sup>  
~~124~~. (New) The method of claim <sup>122</sup>~~123~~, wherein the cell is selected from the group consisting of animal cells, plant cells and microbial cells.

<sup>124</sup>  
~~125~~. (New) The method of claim <sup>122</sup>~~123~~, wherein the first enzyme is a nuclease.

<sup>25</sup>  
~~126~~. (New) The method of claim <sup>122</sup>~~125~~, wherein the nuclease is DNase I.

<sup>126</sup>  
~~127~~. (New) The method of claim <sup>124</sup>~~125~~, wherein the nuclease is a restriction enzyme.

<sup>127</sup>  
~~128~~. (New) The method of claim <sup>122</sup>~~123~~, wherein the second enzyme is a restriction enzyme.

<sup>128</sup>  
~~129~~. (New) The method of claim <sup>127</sup>~~128~~, wherein the restriction enzyme is Sau3A I.

<sup>129</sup>  
~~130~~. (New) The method of claim <sup>128</sup>~~129~~, wherein the second end of the vector molecule is generated by digestion with BamH I.

<sup>130</sup>  
~~131~~. (New) The method of claim <sup>125</sup>~~126~~, wherein, subsequent to step (b), the DNase I ends are converted to blunt ends.

<sup>131</sup>  
~~132~~. (New) The method of claim <sup>130</sup>~~131~~, wherein the first end of the vector molecule is a blunt end.

<sup>132</sup>  
~~133~~. (New) The method of claim <sup>131</sup>~~132~~, wherein the first end of the vector molecule is generated by digestion with EcoRV or SmaI.

<sup>133</sup>  
~~134~~. (New) The method of claim <sup>122</sup>~~123~~ wherein, during steps (b) – (d), the nucleus is embedded in agarose.

<sup>134</sup>  
~~135~~. (New) The method of claim <sup>122</sup>~~123~~, wherein a plurality of different libraries of regulatory DNA sequences are prepared, wherein each library is obtained from a different cell.

<sup>135</sup>  
~~136~~. (New) The method of claim <sup>134</sup>~~135~~ wherein, in step (a), nuclei are obtained from cells at different stages of development.

<sup>136</sup>  
~~137~~. (New) The method of claim <sup>134</sup>~~135~~ wherein, in step (a), nuclei are obtained from cells in different tissues.

<sup>137</sup>  
~~138~~. (New) The method of claim <sup>134</sup>~~135~~ wherein, in step (a), nuclei are obtained from diseased cells and counterpart normal cells.

<sup>138</sup>  
~~139~~. (New) The method of claim <sup>134</sup>~~135~~ wherein, in step (a), nuclei are obtained from infected cells and counterpart uninfected cells.

<sup>139</sup>  
~~140~~. (New) The method of claim <sup>134</sup>~~135~~ wherein, in step (a), nuclei are obtained from cells that express a gene of interest at different levels.

<sup>140</sup>  
~~141~~. (New) The method of claim <sup>122</sup>~~123~~, wherein a plurality of different libraries of regulatory DNA sequences are prepared and, for each library, a different first enzyme is used.

<sup>141</sup>  
~~142~~. (New) The method of claim <sup>140</sup>~~141~~, wherein the different libraries are combined.

<sup>142</sup>  
~~143~~. (New) A method for isolating a collection of polynucleotides comprising cellular regulatory sequences, wherein the method comprises:

(a) contacting cellular chromatin with a probe, wherein the probe reacts with accessible regions of cellular chromatin;

(b) subsequently fragmenting the cellular chromatin to generate a collection of polynucleotide fragments; and

(c) selectively cloning polynucleotide fragments comprising a site of probe reaction.

<sup>143</sup>  
~~144~~ (New) The method of claim <sup>142</sup>~~143~~, wherein reaction of the probe with cellular chromatin results in polynucleotide cleavage at the site of reaction.

<sup>144</sup>  
~~145~~ (New) The method of claim <sup>142</sup>~~143~~, wherein the cellular chromatin is present in an isolated nucleus.

<sup>145</sup>  
~~146~~ (New) The method of claim <sup>144</sup>~~145~~ wherein, in steps (a) and (b), the isolated nucleus is embedded in agarose.

<sup>146</sup>  
~~147~~ (New) The method of claim <sup>142</sup>~~143~~, wherein the probe is an enzyme.

<sup>147</sup>  
~~148~~ (New) The method of claim <sup>146</sup>~~147~~, wherein the enzyme is a nuclease.

<sup>148</sup>  
~~149~~ (New) The method of claim <sup>147</sup>~~148~~, wherein the nuclease is a restriction enzyme.

<sup>149</sup>  
~~150~~ (New) The method of claim <sup>147</sup>~~148~~, wherein the nuclease is DNase I.

<sup>150</sup>  
~~151~~ (New) The method of claim <sup>142</sup>~~143~~ wherein, in step (b), cellular chromatin is fragmented by restriction enzyme digestion.

151

152. (New) The method of claim 150, wherein the restriction enzyme is

Sau3A1.--

B1  
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Attached is a marked-up version of the changes made to the claims by the current amendment. The attached page is captioned "Version with markings to show changes made."